

* * * * * Welcome to STN International * * * * *

<u>NEWS 1</u>		Web Page URLs for STN Seminar Schedule - N. America
<u>NEWS 2</u>		"Ask CAS" for self-help around the clock
<u>NEWS 3</u>	May 12	EXTEND option available in structure searching
<u>NEWS 4</u>	May 12	Polymer links for the POLYLINK command completed in REGISTRY
<u>NEWS 5</u>	May 27	New UPM (Update Code Maximum) field for more efficient patent SDIs in Caplus
<u>NEWS 6</u>	May 27	Caplus super roles and document types searchable in REGISTRY
<u>NEWS 7</u>	Jun 28	Additional enzyme-catalyzed reactions added to CASREACT
<u>NEWS 8</u>	Jun 28	ANTE, AQUALINE, BIOENG, CIVILENG, ENVIROENG, MECHENG, and WATER from CSA now available on STN(R)
<u>NEWS 9</u>	Jul 12	BEILSTEIN enhanced with new display and select options, resulting in a closer connection to BABS
<u>NEWS 10</u>	Jul 30	BEILSTEIN on STN workshop to be held August 24 in conjunction with the 228th ACS National Meeting
<u>NEWS 11</u>	AUG 02	IFIPAT/IFIUDB/IFICDB reloaded with new search and display fields
<u>NEWS 12</u>	AUG 02	Caplus and CA patent records enhanced with European and Japan Patent Office Classifications
<u>NEWS 13</u>	AUG 02	STN User Update to be held August 22 in conjunction with the 228th ACS National Meeting
<u>NEWS 14</u>	AUG 02	The Analysis Edition of STN Express with Discover! (Version 7.01 for Windows) now available
<u>NEWS 15</u>	AUG 04	Pricing for the Save Answers for SciFinder Wizard within STN Express with Discover! will change September 1, 2004
<u>NEWS 16</u>	AUG 27	BIOCOMMERCE: Changes and enhancements to content coverage
<u>NEWS 17</u>	AUG 27	BIOTECHABS/BIOTECHDS: Two new display fields added for legal status data from INPADOC
<u>NEWS 18</u>	SEP 01	INPADOC: New family current-awareness alert (SDI) available
<u>NEWS 19</u>	SEP 01	New pricing for the Save Answers for SciFinder Wizard within STN Express with Discover!
<u>NEWS 20</u>	SEP 01	New display format, HITSTR, available in WPIDS/WPINDEX/WPIX
<u>NEWS 21</u>	SEP 14	STN Patent Forum to be held October 13, 2004, in Iselin, NJ

NEWS EXPRESS JULY 30 CURRENT WINDOWS VERSION IS V7.01, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS INTER General Internet Information

NEWS LOGIN Welcome Banner and News Items

NEWS PHONE Direct Dial and Telecommunication Network Access to STN

NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 10:32:23 ON 21 SEP 2004

=> file req

COST IN U.S. DOLLARS

SINCE FILE
ENTRY

TOTAL
SESSION

h eb c g cg b cg

eb

FULL ESTIMATED COST 0.21 0.21

FILE 'REGISTRY' ENTERED AT 10:32:29 ON 21 SEP 2004
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
 COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file
 provided by InfoChem.

STRUCTURE FILE UPDATES: 20 SEP 2004 HIGHEST RN 748739-98-2
 DICTIONARY FILE UPDATES: 20 SEP 2004 HIGHEST RN 748739-98-2

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when
 conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more
 information enter HELP PROP at an arrow prompt in the file or refer
 to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=>

L1 STRUCTURE UPLOADED

=> 11

L1 IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system.
 For a list of commands available to you in the current file, enter
 "HELP COMMANDS" at an arrow prompt (=>).

=> d 11

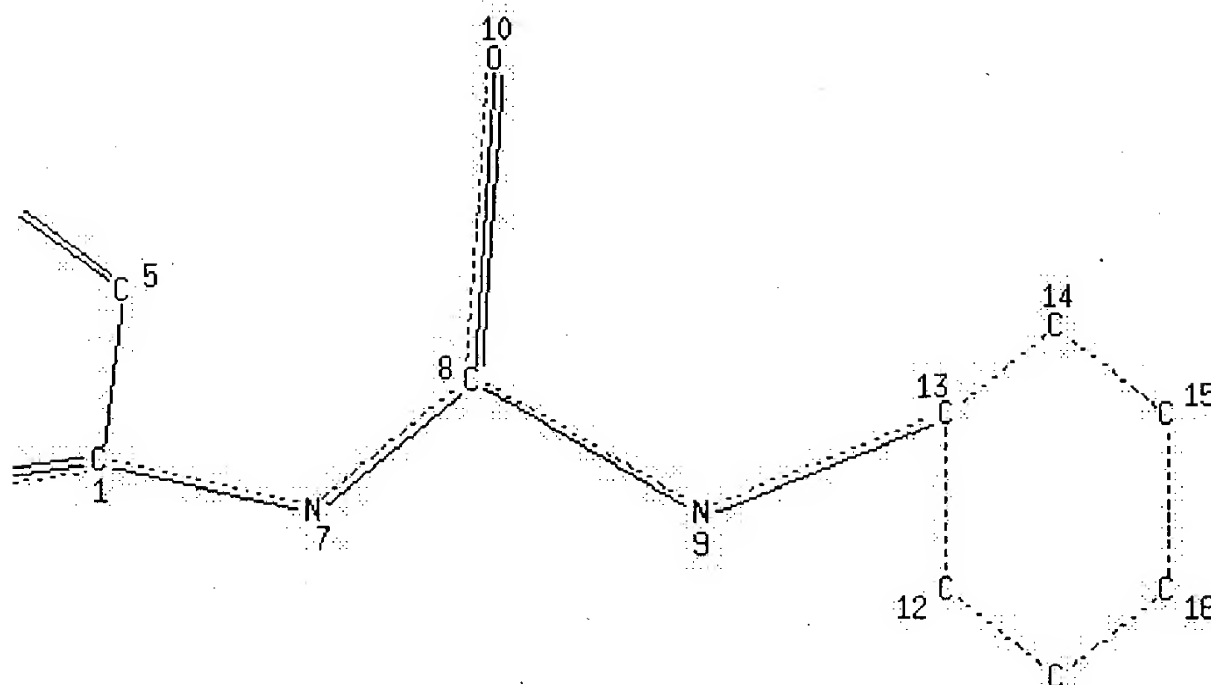
L1 HAS NO ANSWERS

L1 STR

Cb 17Ak 18



Page 1-A



Page 1-B

11

Page 2-B

VAR G1=17/18

NODE ATTRIBUTES:

NSPEC	IS	R	AT	1
NSPEC	IS	R	AT	2
NSPEC	IS	R	AT	3
NSPEC	IS	R	AT	4
NSPEC	IS	R	AT	5
NSPEC	IS	C	AT	6
NSPEC	IS	C	AT	7
NSPEC	IS	C	AT	8
NSPEC	IS	C	AT	9
NSPEC	IS	C	AT	10
NSPEC	IS	R	AT	11
NSPEC	IS	R	AT	12
NSPEC	IS	R	AT	13
NSPEC	IS	R	AT	14
NSPEC	IS	R	AT	15
NSPEC	IS	R	AT	16

DEFAULT MLEVEL IS ATOM

MLEVEL IS CLASS AT 7 8 9 10 17 18

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RSPEC I

NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

$$\Rightarrow \frac{1}{2} \frac{1}{2}$$

SAMPLE SEARCH INITIATED 10:34:39 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 53 TO ITERATE

100.0% PROCESSED 53 ITERATIONS

23 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH ** COMPLETE**

PROJECTED ITERATIONS: 624 TO 1496

h eb c g cg b cg

eb

PROJECTED ANSWERS: 173 TO 747

L2 23 SEA SSS SAM L1

=> s 11 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS

DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END:y

FULL SEARCH INITIATED 10:34:43 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 893 TO ITERATE

100.0% PROCESSED 893 ITERATIONS

377 ANSWERS

SEARCH TIME: 00.00.01

L3 377 SEA SSS FUL L1

=> file hcaplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

156.68

156.89

FILE 'HCAPLUS' ENTERED AT 10:34:46 ON 21 SEP 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 21 Sep 2004 VOL 141 ISS 13

FILE LAST UPDATED: 20 Sep 2004 (20040920/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13

L4 34 L3

=> file reg

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

7.08

163.97

FILE 'REGISTRY' ENTERED AT 10:36:32 ON 21 SEP 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 20 SEP 2004 HIGHEST RN 748739-98-2

DICTIONARY FILE UPDATES: 20 SEP 2004 HIGHEST RN 748739-98-2

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more
information enter HELP PROP at an arrow prompt in the file or refer
to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=>

L5 STRUCTURE UPLOADED

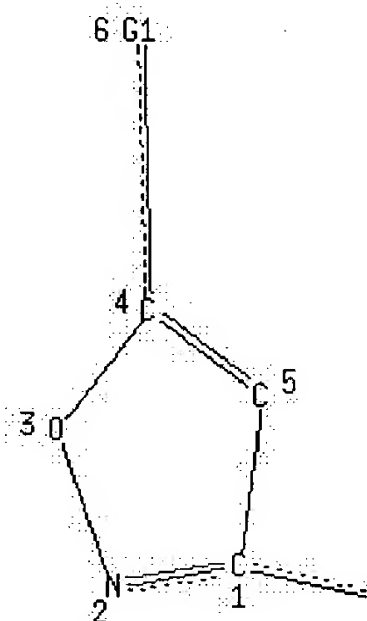
=> d 15

L5 HAS NO ANSWERS

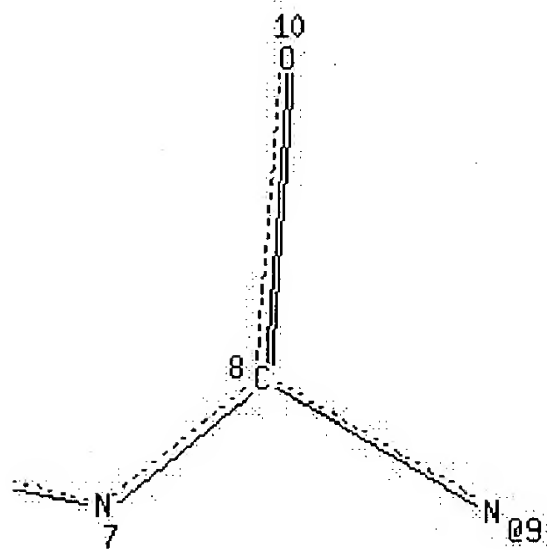
L5 STR

0 21 5 22

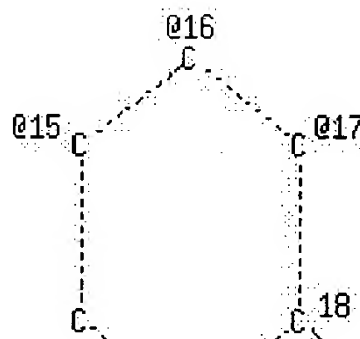
Cb 19Ak 20



Page 1-A



Page 1-B



12
C4
Page 1-C



Page 2-B
VAR G1=19/20
VAR G2=21/22
VPA 9-13/14/15/16/17 S
NODE ATTRIBUTES:
NSPEC IS R AT 1
NSPEC IS R AT 2
NSPEC IS R AT 3
NSPEC IS R AT 4
NSPEC IS R AT 5
NSPEC IS C AT 6
NSPEC IS C AT 7
NSPEC IS C AT 8
NSPEC IS C AT 9
NSPEC IS C AT 10
NSPEC IS C AT 11
NSPEC IS C AT 12
NSPEC IS R AT 13
NSPEC IS R AT 14
NSPEC IS R AT 15
NSPEC IS R AT 16
NSPEC IS R AT 17
NSPEC IS R AT 18
DEFAULT MLEVEL IS ATOM
MLEVEL IS CLASS AT 7 8 9 10 19 20 21 22
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RSPEC 13
NUMBER OF NODES IS 22

STEREO ATTRIBUTES: NONE

=> s 15
SAMPLE SEARCH INITIATED 10:39:48 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 19 TO ITERATE

100.0% PROCESSED 19 ITERATIONS 8 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 119 TO 641
PROJECTED ANSWERS: 8 TO 329

L6 8 SEA SSS SAM L5

=> s 15 full

THE ESTIMATED SEARCH COST FOR FILE 'REGISTRY' IS 155.00 U.S. DOLLARS
 DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N or END:Y
 FULL SEARCH INITIATED 10:39:52 FILE 'REGISTRY'
 FULL SCREEN SEARCH COMPLETED - 329 TO ITERATE

100.0% PROCESSED 329 ITERATIONS 189 ANSWERS
 SEARCH TIME: 00.00.01

L7 189 SEA SSS FUL L5

=> file hcaplus

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
157.52	321.49

FULL ESTIMATED COST

FILE 'HCAPLUS' ENTERED AT 10:39:55 ON 21 SEP 2004
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
 COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 21 Sep 2004 VOL 141 ISS 13
 FILE LAST UPDATED: 20 Sep 2004 (20040920/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l7

L8 10 L7

=> s l8 and dumas, j?/au

677 DUMAS, J?/AU

L9 3 L8 AND DUMAS, J?/AU

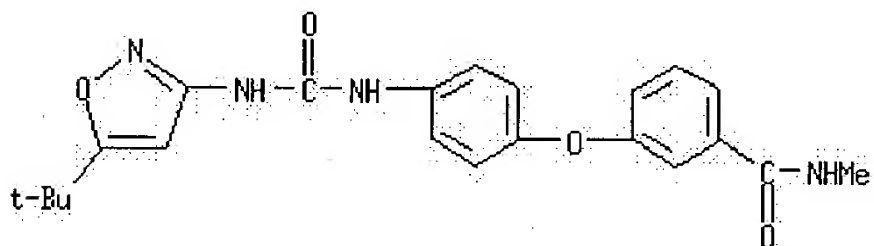
=> d l9, ibib abs fhitstr, 1-3

L9 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

Full Text References

ACCESSION NUMBER: 2004:51821 HCAPLUS
 DOCUMENT NUMBER: 140:296858
 TITLE: Omega-carboxypyridyl substituted ureas as Raf kinase inhibitors: SAR of the amide substituent
 AUTHOR(S): Khire, Uday R.; Bankston, Donald; Barbosa, James; Brittelli, David R.; Caringal, Yolanda; Carlson, Robert; Dumas, Jacques; Gane, Todd; Heald, Sarah L.; Hibner, Barbara; Johnson, Jeffrey S.; Katz, Michael E.; Kennure, Nancy; Kingery-Wood, Jill; Lee, Wendy; Liu, Xiao-Gao; Lowinger, Timothy B.; McAlexander, Ian;

Monahan, Mary-Katherine; Natero, Reina; Renick, Joel;
 Riedl, Bernd; Rong, Hong; Sibley, Robert N.; Smith,
 Roger A.; Wolanin, Donald
 CORPORATE SOURCE: Department of Chemistry Research, Bayer Research
 Center, West Haven, CT, 06516, USA
 SOURCE: Bioorganic & Medicinal Chemistry Letters (2004),
 14(3), 783-786
 CODEN: BMCLE8; ISSN: 0960-894X
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Bis-aryl ureas have been disclosed previously as a potent class of Raf
 kinase inhibitors. Modifications in the amide portion led to an
 improvement in aq. soly., an important characteristic for an oral drug.
 Based on this finding, we hypothesize that this portion of the mol. is
 directed towards the solvent in Raf-1.
 IT 228999-58-4
 RL: PAC (Pharmacological activity); BIOL (Biological study)
 (structure and Raf kinase inhibitor activity of amide substituent of
 omega-carboxypyridyl substituted ureas)
 RN 228999-58-4 HCAPLUS
 CN Benzamide, 3-[4-[[[5-(1,1-dimethylethyl)-3-isoxazolyl]amino]carbonyl]amin
 o]phenoxy]-N-methyl- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

Full
Text
References

ACCESSION NUMBER: 1999:425745 HCAPLUS
 DOCUMENT NUMBER: 131:87909
 TITLE: Inhibition of p38 kinase activity using substituted
 heterocyclic ureas
 INVENTOR(S): **Dumas, Jacques**; Khire, Uday; Lowinger, Timothy
 Bruno; Paulsen, Holger; Riedl, Bernd; Scott, William
 J.; Smith, Roger A.; Wood, Jill E.; Hatoum-Mokdad,
 Holia; Johnson, Jeffrey; Lee, Wendy; Redman, Aniko
 PATENT ASSIGNEE(S): Bayer Corporation, USA
 SOURCE: PCT Int. Appl., 126 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9932111	A1	19990701	WO 1998-US26080	19981222
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,				

MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
 TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
 FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
 CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

CA 2315720 AA 19990701 CA 1998-2315720 19981222

AU 9919971 A1 19990712 AU 1999-19971 19981222

AU 739642 B2 20011018

EP 1041982 A1 20001011 EP 1998-964709 19981222

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO

JP 2001526223 T2 20011218 JP 2000-525102 19981222

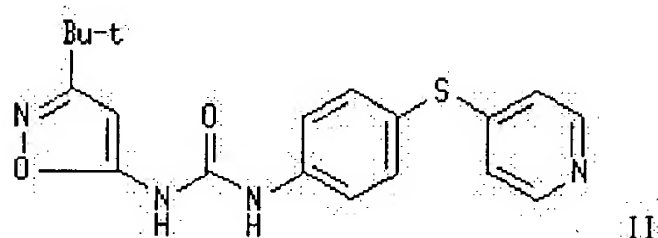
PRIORITY APPLN. INFO.:

US 1997-995750 A 19971222

WO 1998-US26080 W 19981222

OTHER SOURCE(S): MARPAT 131:87909

GI



AB A method for treatment of p38-mediated disease other than cancer comprises administration of ANHCONHB [I; A = substituted isoxazolyl, pyrazolyl, thienyl, furyl; B = (substituted) mono-, di-, or tricyclic aryl, heteroaryl contg. ≥ 1 5-6 membered arom. structure contg. 0-4 N, O, or S atoms]. Reaction of 4-(4-pyridinylthio)aniline with 3-tert-butyl-5-isoxazolyl isocyanate in toluene gave title compd. II. In an in vitro p38 kinase assay, I displayed IC50 values of 1-10 μ M.

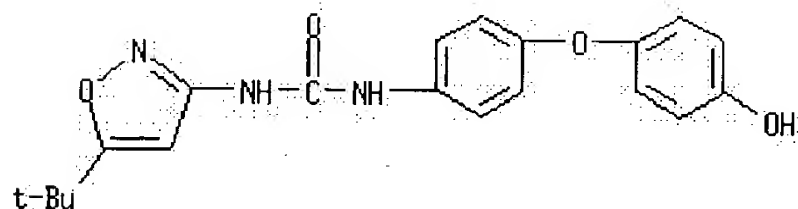
IT **228999-08-4P**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(prepn. of substituted heterocyclic ureas for treatment of p38 kinase-mediated diseases other than cancer)

RN **228999-08-4** HCAPLUS

CN Urea, N-[5-(1,1-dimethylethyl)-3-isoxazolyl]-N'-[4-(4-hydroxyphenoxy)phenyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

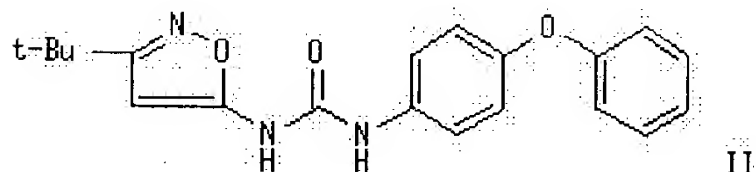
Full Text References

ACCESSION NUMBER: 1999:425740 HCAPLUS

DOCUMENT NUMBER: 131:73648
 TITLE: Inhibition of raf kinase using substituted heterocyclic ureas
 INVENTOR(S): **Dumas, Jacques**; Khire, Uday; Lowinger, Timothy Bruno; Paulsen, Holger; Riedl, Bernd; Scott, William J.; Smith, Roger A.; Wood, Jill E.; Hatoum-Mokdad, Holia; Johnson, Jeffrey; Lee, Wendy; Redman, Aniko
 PATENT ASSIGNEE(S): Bayer Corporation, USA
 SOURCE: PCT Int. Appl., 163 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9932106	A1	19990701	WO 1998-US26078	19981222
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2315717	AA	19990701	CA 1998-2315717	19981222
AU 9921989	A1	19990712	AU 1999-21989	19981222
EP 1047418	A1	20001102	EP 1998-965981	19981222
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
TR 200002618	T2	20010420	TR 2000-200002618	19981222
JP 2001526220	T2	20011218	JP 2000-525097	19981222
BR 9814374	A	20020514	BR 1998-14374	19981222
RU 2232015	C2	20040710	RU 2000-120184	19981222
NO 2000003232	A	20000821	NO 2000-3232	20000621
BG 104597	A	20010228	BG 2000-104597	20000712
PRIORITY APPLN. INFO.:			US 1997-996343	A 19971222
			WO 1998-US26078	W 19981222

OTHER SOURCE(S): MARPAT 131:73648
 GI



AB A method for treatment of cancerous cell growth mediated by raf kinase comprises administration of urea derivs. ANHCONHB [I; A = substituted isoxazoly, thienyl, thiadiazoly, furyl, pyrazoly, etc.; B = (substituted) mono-, di-, or tricyclic aryl, heteroaryl contg. ≥ 1 5-6 membered arom. structure contg. 0-4 N, O, or S atoms]. Reaction of 4-phenyloxyphenyl isocyanate with 5-amino-3-tert-butylisoxazole in methylene chloride and heating at reflux temp. for 2 days gave title compd. II. In an in vitro raf kinase assay, I displayed IC₅₀ values of 1-10 μ M.

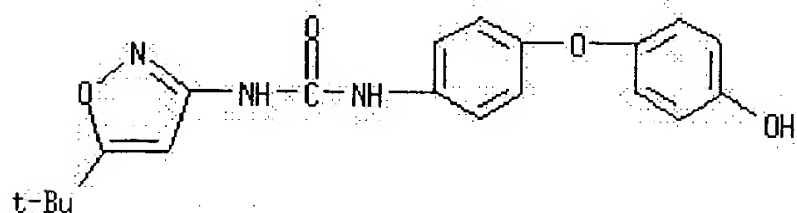
IT 228999-08-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(prepn. of substituted heterocyclic ureas for treatment of cancerous cell growth mediated by raf kinase)

RN 228999-08-4 HCAPLUS

CN Urea, N-[5-(1,1-dimethylethyl)-3-isoxazolyl]-N'-[4-(4-hydroxyphenoxy)phenyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 10:32:23 ON 21 SEP 2004)

FILE 'REGISTRY' ENTERED AT 10:32:29 ON 21 SEP 2004

L1 STRUCTURE UPLOADED

L2 23 S L1

L3 377 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 10:34:46 ON 21 SEP 2004

L4 34 S L3

FILE 'REGISTRY' ENTERED AT 10:36:32 ON 21 SEP 2004

L5 STRUCTURE UPLOADED

L6 8 S L5

L7 189 S L5 FULL

FILE 'HCAPLUS' ENTERED AT 10:39:55 ON 21 SEP 2004

L8 10 S L7

L9 3 S L8 AND DUMAS, J?/AU

=> s l8 not l9

L10 7 L8 NOT L9

=> s l10 and khire, u?/au

41 KHIRE, U?/AU

L11 0 L10 AND KHIRE, U?/AU

=> s l10 and lowinger, t?/au

44 LOWINGER, T?/AU

L12 1 L10 AND LOWINGER, T?/AU

=> d l12, ibib abs thitstr, 1

L12 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN

FULL
Text References

ACCESSION NUMBER: 2001:746592 HCAPLUS

h eb c g cg b cg

eb

DOCUMENT NUMBER: 136:95577

TITLE: Discovery of heterocyclic ureas as a new class of raf kinase inhibitors: identification of a second generation lead by a combinatorial chemistry approach

AUTHOR(S): Smith, R. A.; Barbosa, J.; Blum, C. L.; Bobko, M. A.; Caringal, Y. V.; Dally, R.; Johnson, J. S.; Katz, M. E.; Kennure, N.; Kingery-Wood, J.; Lee, W.; **Lowinger, T. B.**; Lyons, J.; Marsh, V.; Rogers, D. H.; Swartz, S.; Walling, T.; Wild, H.

CORPORATE SOURCE: Department of Chemistry Research, Bayer Research Center, West Haven, CT, 06516, USA

SOURCE: Bioorganic & Medicinal Chemistry Letters (2001), 11(20), 2775-2778
CODEN: BMCLE8; ISSN: 0960-894X

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

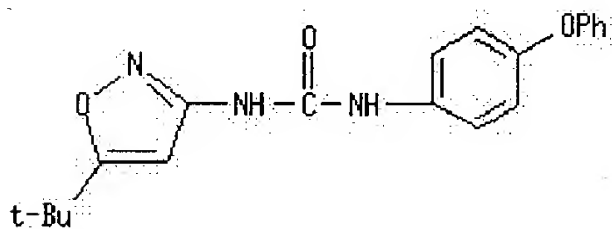
LANGUAGE: English

AB Heterocyclic ureas, such as N-3-thienyl N'-aryl ureas, have been identified as novel inhibitors of raf kinase, a key mediator in the ras signal transduction pathway. Structure-activity relationships were established, and the potency of the screening hit was improved 10-fold to IC₅₀=1.7 µM. A combinatorial synthesis approach enabled the identification of a breakthrough lead (IC₅₀=0.54 µM) for a second generation series of heterocyclic urea raf kinase inhibitors.

IT 228998-90-1P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(heterocyclic ureas as raf kinase inhibitors)

RN 228998-90-1 HCAPLUS

CN Urea, N-[5-(1,1-dimethylethyl)-3-isoxazoly]-N'-(4-phenoxyphenyl)- (9CI)
(CA INDEX NAME)



REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 10:32:23 ON 21 SEP 2004)

FILE 'REGISTRY' ENTERED AT 10:32:29 ON 21 SEP 2004

L1 STRUCTURE UPLOADED
L2 23 S L1
L3 377 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 10:34:46 ON 21 SEP 2004

L4 34 S L3

FILE 'REGISTRY' ENTERED AT 10:36:32 ON 21 SEP 2004

L5 STRUCTURE UPLOADED
L6 8 S L5

L7 189 S L5 FULL

FILE 'HCAPLUS' ENTERED AT 10:39:55 ON 21 SEP 2004

L8 10 S L7

L9 3 S L8 AND DUMAS, J?/AU

L10 7 S L8 NOT L9

L11 0 S L10 AND KHIRE, U?/AU

L12 1 S L10 AND LOWINGER, T?/AU

=> s l10 not l12

L13 6 L10 NOT L12

=> s l13 and paulsen, h?/au

670 PAULSEN, H?/AU

L14 0 L13 AND PAULSEN, H?/AU

=> s l13 and riedl, b?/au

167 RIEDL, B?/AU

L15 0 L13 AND RIEDL, B?/AU

=> s l13 and scott, w?/au

1980 SCOTT, W?/AU

L16 0 L13 AND SCOTT, W?/AU

=> s l13 and smith, r?/au

13624 SMITH, R?/AU

L17 0 L13 AND SMITH, R?/AU

=> s l13 and wood, j?/au

3934 WOOD, J?/AU

L18 0 L13 AND WOOD, J?/AU

=> s l13 and hatoum-mokdad, h?/au

27 HATOUM-MOKDAD, H?/AU

L19 0 L13 AND HATOUM-MOKDAD, H?/AU

=> s l13 and johnson, j?/au

7801 JOHNSON, J?/AU

L20 0 L13 AND JOHNSON, J?/AU

=> s l13 and lee, w?/au

8975 LEE, W?/AU

L21 0 L13 AND LEE, W?/AU

=> s l13 and redman, a?/au

33 REDMAN, A?/AU

L22 0 L13 AND REDMAN, A?/AU

=> s l13 and sibley, r?/au

193 SIBLEY, R?/AU

L23 0 L13 AND SIBLEY, R?/AU

=> s l13 and renick, f?/au

0 RENICK, F?/AU

L24 0 L13 AND RENICK, F?/AU

=> s l13 and renick, j?/au

14 RENICK, J?/AU

L25 0 L13 AND RENICK, J?/AU

=> d his

(FILE 'HOME' ENTERED AT 10:32:23 ON 21 SEP 2004)

FILE 'REGISTRY' ENTERED AT 10:32:29 ON 21 SEP 2004

L1 STRUCTURE UPLOADED
 L2 23 S L1
 L3 377 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 10:34:46 ON 21 SEP 2004

L4 34 S L3

FILE 'REGISTRY' ENTERED AT 10:36:32 ON 21 SEP 2004

L5 STRUCTURE UPLOADED
 L6 8 S L5
 L7 189 S L5 FULL

FILE 'HCAPLUS' ENTERED AT 10:39:55 ON 21 SEP 2004

L8 10 S L7
 L9 3 S L8 AND DUMAS, J?/AU
 L10 7 S L8 NOT L9
 L11 0 S L10 AND KHIRE, U?/AU
 L12 1 S L10 AND LOWINGER, T?/AU
 L13 6 S L10 NOT L12
 L14 0 S L13 AND PAULSEN, H?/AU
 L15 0 S L13 AND RIEDL, B?/AU
 L16 0 S L13 AND SCOTT, W?/AU
 L17 0 S L13 AND SMITH, R?/AU
 L18 0 S L13 AND WOOD, J?/AU
 L19 0 S L13 AND HATOUM-MOKDAD, H?/AU
 L20 0 S L13 AND JOHNSON, J?/AU
 L21 0 S L13 AND LEE, W?/AU
 L22 0 S L13 AND REDMAN, A?/AU
 L23 0 S L13 AND SIBLEY, R?/AU
 L24 0 S L13 AND RENICK, F?/AU
 L25 0 S L13 AND RENICK, J?/AU

=> d l13, ibib abs fhitr, 1-6

L13 ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

Full Text
 L13 Reference

ACCESSION NUMBER: 2004:589418 HCAPLUS
 DOCUMENT NUMBER: 141:117198
 TITLE: Therapeutic agent for wet age-related macular degeneration
 INVENTOR(S): Matsuno, Kiyoshi; Koyama, Shinji
 PATENT ASSIGNEE(S): Santen Pharmaceutical Co., Ltd., Japan; Kirin Beer Kabushiki Kaisha
 SOURCE: PCT Int. Appl., 21 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004060373	A1	20040722	WO 2003-JP16854	20031226
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,				

CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
 GE, GH, GM, HR, HU, ID, IL, IN, IS, KE, KG, KP, KR, KZ, LC, LK,
 LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
 OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
 TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ,
 BY, KG, KZ, MD
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
 BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU,
 MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
 GQ, GW, ML, MR, NE, SN, TD, TG

JP 2004217649

A2

20040805

JP 2003-431849

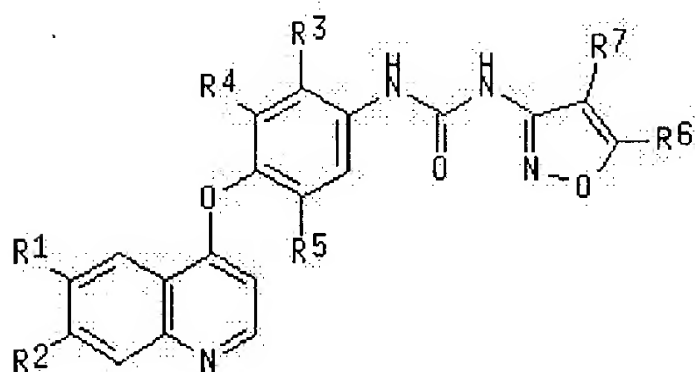
20031226

PRIORITY APPLN. INFO.:

JP 2002-379857

A 20021227

GI



I

AB A therapeutic agent for wet age-related macular degeneration which contains as an active ingredient an N-quinolyloxyphenyl-N'-isoxazolyurea deriv. represented by the general formula (I; wherein R1 and R2 each is C1-6 alkoxy; R3 is halogeno; R4 and R5 each is hydrogen, halogeno, etc.; and R6 and R7 each is hydrogen, halogeno, C1-4 alkyl, etc.). The compd. has excellent choroidal angiogenesis inhibitory activity and is useful in treatments for wet age-related macular degeneration.

IT 475108-18-0, N-{2-Chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl}-N'-(5-methyl-3-isoxazolyl)urea

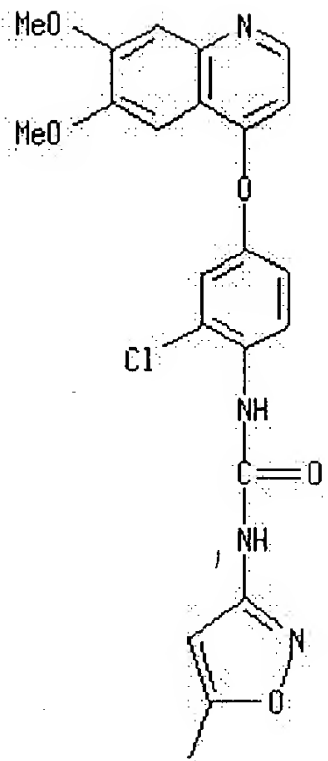
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(N-quinolyloxyphenyl-N'-isoxazolyurea derivs. as therapeutic agents for wet age-related macular degeneration)

RN 475108-18-0 HCAPLUS

CN Urea, N-[2-chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

Me

L13 ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

Full Text References

ACCESSION NUMBER: 2004:354935 HCAPLUS
DOCUMENT NUMBER: 140:363009
TITLE: N-[2-Chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)urea salt crystals
INVENTOR(S): Matsunaga, Naoki; Yoshida, Satoshi; Yoshino, Ayako; Nakajima, Tatsuo
PATENT ASSIGNEE(S): Kirin Beer Kabushiki Kaisha, Japan
SOURCE: PCT Int. Appl., 115 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004035572	A1	20040429	WO 2003-JP13439	20031021
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			JP 2002-306101	A 20021021

AB This invention provides crystals of pharmaceutically acceptable salts of N-[2-chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)urea. The salt crystals are used in treating a disease selected from the group consisting of tumor, diabetic retinopathy, rheumatoid arthritis, psoriasis, atheroma arteriosclerosis, Kaposi's sarcoma and exudative age-related macular degeneration. The salt crystals have properties appropriate for preps. for oral administration.

IT 682745-43-3P

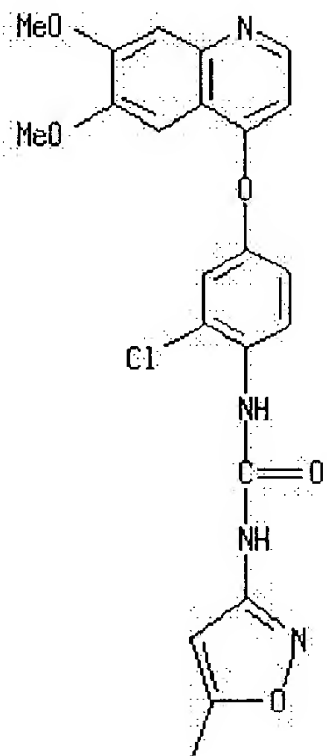
RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(N-[2-Chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)urea salt crystals with activity against tumor, diabetic retinopathy, rheumatoid arthritis, psoriasis, neovascularization, etc.)

RN 682745-43-3 HCAPLUS

CN Urea, N-[2-chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)-, monohydrochloride (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

HCl

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 3 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

Full Text ☐
References ☐

ACCESSION NUMBER: 2004:182368 HCAPLUS
DOCUMENT NUMBER: 140:229401
TITLE: Three hybrid assay system for isolating ligand-binding polypeptides and for isolating small mol. ligands
INVENTOR(S): Come, Jon H.; Becker, Frank; Kley, Nikolai A.; Reichel, Christoph

PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 238 pp., Cont.-in-part of U.S. Ser. No. 91,177.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004043388	A1	20040304	US 2002-234985	20020903
US 2003165873	A1	20030904	US 2002-91177	20020304
PRIORITY APPLN. INFO.:			US 2001-272932P	P 20010302
			US 2001-278233P	P 20010323
			US 2001-329437P	P 20011015
			US 2002-91177	A2 20020304

AB The invention provides compns. and methods for isolating ligand-binding polypeptides for a user-specified ligand, and for isolating small mol. ligands for a user-specified target polypeptide using an improved class of hybrid ligand compds. Prepn. of compds., e.g a methotrexate moiety linked by a polyethylene glycol moiety to dexamethasone, is described.

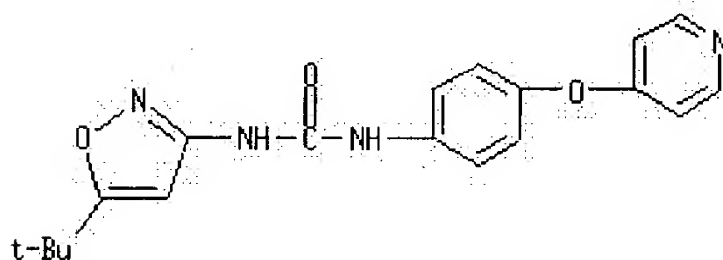
IT **228999-48-2D**, conjugates

RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(three hybrid assay system for isolating ligand-binding polypeptides and for isolating small mol. ligands)

RN **228999-48-2** HCAPLUS

CN Urea, N-[5-(1,1-dimethylethyl)-3-isoxazolyl]-N'-[4-(4-pyridinyloxy)phenyl]-(9CI) (CA INDEX NAME)



L13 ANSWER 4 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

Full Text **SHIRI** References

ACCESSION NUMBER: 2002:849617 HCAPLUS
 DOCUMENT NUMBER: 137:370101
 TITLE: Preparation of quinoline derivatives having azolyl group and quinazoline derivatives as antitumor agents
 INVENTOR(S): Kubo, Kazuo; Sakai, Teruyuki; Nagao, Rika; Fujiwara, Yasunari; Isoe, Toshiyuki; Hasegawa, Kazumasa
 PATENT ASSIGNEE(S): Kirin Beer Kabushiki Kaisha, Japan
 SOURCE: PCT Int. Appl., 89 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----

WO 2002088110 A1 20021107 WO 2002-JP4279 20020426

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

JP 2003012668 A2 20030115 JP 2002-126869 20020426

US 2003087907 A1 20030508 US 2002-132473 20020426

EP 1382604 A1 20040121 EP 2002-724651 20020426

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

BR 2002009216 A 20040706 BR 2002-9216 20020426

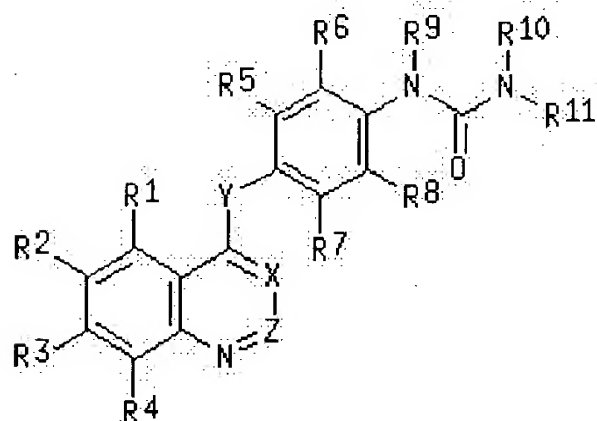
NO 2003004595 A 20031219 NO 2003-4595 20031014

JP 2004224800 A2 20040812 JP 2004-101164 20040330

PRIORITY APPLN. INFO.: JP 2001-132775 A 20010427

OTHER SOURCE(S): MARPAT 137:370101 JP 2002-126869 A3 20020426

GI WO 2002-JP4279 W 20020426



AB N-[(4-quinolinyl or 4-quinazolinyl)thio or -oxy]phenyl-N'-azolylylurea derivs. represented by the formula (I) or pharmaceutically acceptable salts or solvates thereof [wherein X, Z = CH, N; Y = O, S; R1, R2, R3 = H, NO2, NH2, each (un)substituted C1-6 alkyl or alkoxy or C2-6 alkenyl or alkynyl; R4 = H; R5-R8 = H, halo, C1-4 alkyl, alkoxy, or alkylthio, CF3, NO2, NH2; R9, R10 = C1-6 alkyl, each (un)substituted C1-4 alkylcarbonyl or C1-6 alkyl; R11 = (un)substituted azolyl] are prepd. These compds. are useful for the treatment of tumor, diabetic retinopathy, chronic articular rheumatism, psoriasis, atherosclerosis, and Kaposi's sarcoma. They are also used for inhibiting neovascularization of a target blood vessel by contacting them with vascular endothelial cells of the target blood vessel. Thus, 100 mg 2-chloro-4-[(6,7-dimethoxy-4-quinazolinyl)oxy]aniline was dissolved in 5 mL CHCl3 and 0.5 mL Et3N, treated with a soln. of 100 mg triphosgene in CHCl3, and stirred at room temp. for 15 min, followed by adding 49 mg 2-aminothiazole, and the resulting mixt. was stirred at room temp. overnight to give 31 mg N-[2-chloro-4-[(6,7-dimethoxy-4-quinazolinyl)oxy]phenyl]-N'-(1,3-thiazol-2-yl)urea (II). II at 20 mg/kg/day for 9 days inhibited the growth of human lung cancer transplanted in nude mice by 92.0%. The compds. I in vitro showed IC50 of 0.001-0.0697 μ M for inhibiting the phosphorylation of

the intracellular domain of human vascular endothelial cell growth factor (VEGF) receptor KDR (kinase insert domain-contg. receptor) in IH3T3 cell expressing human KDR.

IT **475108-18-0P**, N-[2-Chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)urea

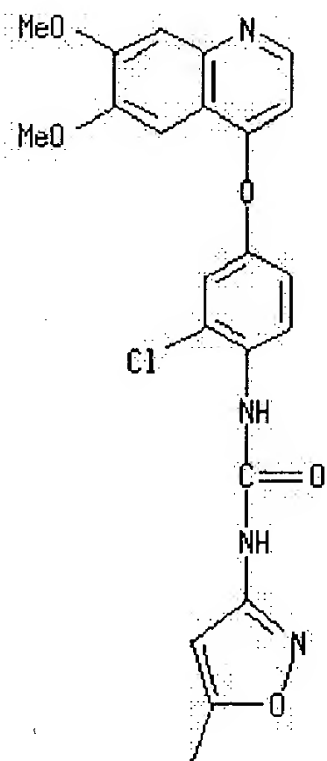
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of N-[(4-quinolinyl or 4-quinazolinyl)oxy]phenyl-N'-azolylurea derivs. as neovascularization inhibitors for treatment of tumor, diabetic retinopathy, chronic articular rheumatism, psoriasis, atherosclerosis, and Kaposi's sarcoma)

RN **475108-18-0** HCAPLUS

CN Urea, N-[2-chloro-4-[(6,7-dimethoxy-4-quinolyl)oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

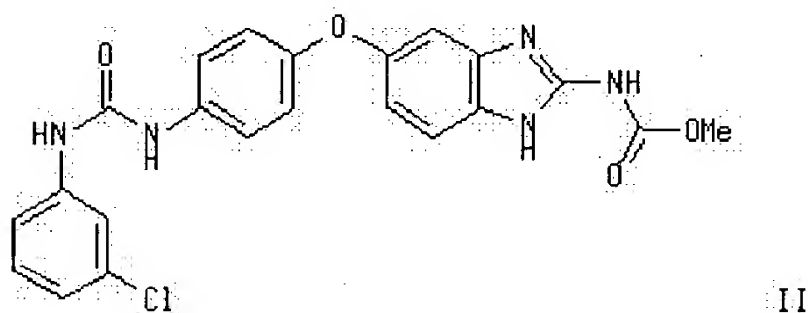
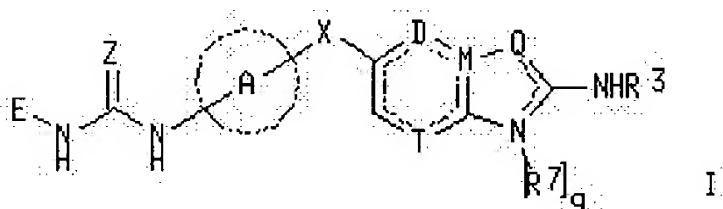
Full Text References

ACCESSION NUMBER: 2002:428885 HCAPLUS
DOCUMENT NUMBER: 137:6179
TITLE: Preparation of benzimidazoles as TIE-2 and/or VEGFR2 inhibitors
INVENTOR(S): Cheung, Mui; Harris, Philip Anthony; Hasegawa, Masaichi; Ida, Satoru; Kano, Kazuya; Nishigaki, Naohiko; Sato, Hideyuki; Veal, James Martin; Washio, Yoshiaki; West, Rob I.
PATENT ASSIGNEE(S): Glaxo Group Limited, UK; Glaxosmithkline K.K.
SOURCE: PCT Int. Appl., 217 pp.

CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002044156	A2	20020606	WO 2001-US44553	20011128
WO 2002044156	A3	20021017		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2002032439	A5	20020611	AU 2002-32439	20011128
EP 1341771	A2	20030910	EP 2001-991963	20011128
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
JP 2004517080	T2	20040610	JP 2002-546526	20011128
US 2004082583	A1	20040429	US 2003-433128	20031112
PRIORITY APPLN. INFO.:				
			US 2000-253868P	P 20001129
			US 2001-310939P	P 20010808
			WO 2001-US44553	W 20011128

OTHER SOURCE(S): MARPAT 137:6179
 GI



AB The title compds. [I; E = (un)substituted aryl, heteroaryl; A = aryl, heteroaryl, heterocyclyl; X = S, O, SO₂, SO, CH₂, CHOH, CO; Z = O, S; p = 0-1; q = 0-1; D = CH, T = CR₈, M = C and Q = NT_{7p}, wherein p = 0 and q = 1; or D = CH, T = CR₈, M = C and Q = NR_{7p}, wherein p = 1 and q = 0, or D = CH, T = CR₈, M = C and Q = S or O, wherein q = 0; or D = N, T = CR₈, M = C and Q = NR_{7p}, wherein either p or q = 0 and the other = 1; or D = CH, T = N, M = C and Q = NR_{7p}, wherein either p or q = 0 and the other = 1; or D = CH, T = CR₈, M = N and Q = CH, wherein q = 0; R₁ = alkyl, haloalkyl, aryl, etc.; R₂ = H, alkyl, aryl, etc.; R₃ = alkylene or alkylene substituted by oxo, and is linked together with N atom to which it is attached and to one

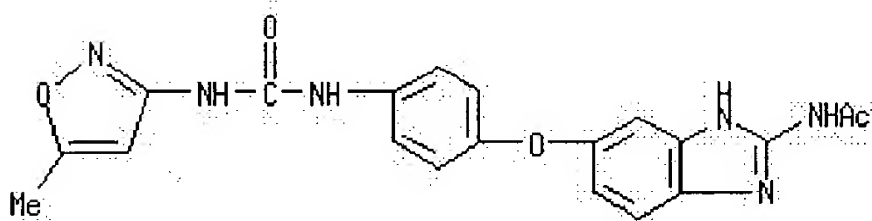
of the benzimidazole N atoms to form a heterocyclic compd. fused to the benzimidazole; R7 = H, alkyl, etc.; R8 = H, halo] and their salts, useful in the treatment of hyperproliferative diseases, were prepd. Thus, reacting Me [5-(4-aminophenoxy)-1H-benzimidazol-2-yl]carbamate (prepn. given) with 3-chlorophenyl isocyanate in THF afforded 69% II which showed pIC50 of > 7.0 in TIE-2 and VEGFR2 enzyme assays.

IT **433225-41-3P**

RL: CPN (Combinatorial preparation); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); CMBI (Combinatorial study); PREP (Preparation); USES (Uses)
(prepn. of benzimidazoles as TIE-2 and/or VEGFR2 inhibitors)

RN **433225-41-3 HCAPLUS**

CN Acetamide, N-[5-[4-[[[(5-methyl-3-isoxazolyl)amino]carbonyl]amino]phenoxy]-1H-benzimidazol-2-yl]- (9CI) (CA INDEX NAME)



L13 ANSWER 6 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

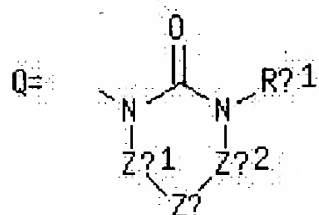
Full Text References

ACCESSION NUMBER: 2002:314913 HCAPLUS
DOCUMENT NUMBER: 136:340689
TITLE: Preparation of urea derivatives containing nitrogenous aromatic ring compounds as inhibitors of angiogenesis
INVENTOR(S): Funahashi, Yasuhiro; Tsuruoka, Akihiko; Matsukura, Masayuki; Haneda, Toru; Fukuda, Yoshio; Kamata, Junichi; Takahashi, Keiko; Matsushima, Tomohiro; Miyazaki, Kazuki; Nomoto, Kenichi; Watanabe, Tatsuo; Obaishi, Hiroshi; Yamaguchi, Atsumi; Suzuki, Sachi; Nakamura, Katsuji; Mimura, Fusayo; Yamamoto, Yuji; Matsui, Junji; Matsui, Kenji; Yoshida, Takako; Suzuki, Yasuyuki; Arimoto, Itaru
PATENT ASSIGNEE(S): Eisai Co., Ltd., Japan
SOURCE: PCT Int. Appl., 699 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002032872	A1	20020425	WO 2001-JP9221	20011019
WO 2002032872	C1	20020926		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2001095986	A5	20020429	AU 2001-95986	20011019

EP 1415987	A1	20040506	EP 2001-976786	20011019
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY, TR				
NO 2003001731	A	20030619	NO 2003-1731	20030414
US 2004053908	A1	20040318	US 2003-420466	20030418
PRIORITY APPLN. INFO.:			JP 2000-320420	A 20001020
			JP 2000-386195	A 20001220
			JP 2001-46685	A 20010222
			WO 2001-JP9221	W 20011019

OTHER SOURCE(S): *ub* MARPAT 136:340689
GI



AB N-aryl or N-heteroarylurea derivs. represented by the general formula Ag-Xg-Yg-Tg1 or salts thereof, or hydrates of both [wherein Ag = (un)substituted C6-14 aryl or 5- to 14-membered heterocyclic group; Xg = single bond, O, S, C1-6 alkylene, SO, SO₂, (un)substituted NH; Yg = (un)substituted C6-14 aryl, 5- to 14-membered heterocyclic group, C1-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl-C1-6 alkyl, 5- to 14-membered heteroaryl-C1-6 alkyl, (CH₂)_gSO₂ (g = 1-8), (CH₂)_{fa}CH:CH(CH₂)_{fb} (fa, fb = 0, 1, 2, 3), etc.; and Tg1 = a group of the general formula -Eg-CO-NRg1(Zg) or Q; wherein Eg = a single bond, (un)substituted NH; Rg1 = H, (un)substituted C1-6 alkyl, C2-6 alkenyl, C2-6 alkynyl, C3-8 aliph. hydrocarbyl, etc.; Zg = C1-8 alkyl, C3-8 alicyclic hydrocarbyl, C6-14 aryl, etc.; Zg1, Zg2 = (a) a single bond, (b) C1-6 alkylene optionally having ≥1 atoms selected from O, S, and N in the middle or the terminus of the chain and optionally substituted with oxo, (c) (un)substituted C2-6 alkenyl] are prepd. These compds. are also inhibitors of vascular endothelial growth factor receptor kinase (VEGFR2 kinase) and are useful as antitumor agents against hemangioma, pancreatic cancer, stomach cancer, colon cancer, breast cancer, prostate cancer, lung cancer, brain tumor, leukemia, or ovarian cancer, as cancer metastasis inhibitors, and for the treatment of retina neovascularization, diabetic retinopathy, atherosclerosis, or inflammatory diseases such as osteoarthritis, rheumatoid arthritis, psoriasis, or delayed hypersensitivity. Thus, to soln. of 334 mg 4-[6-(4-benzyloxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenylamine in 4 mL DMF were added 0.066 mL pyridine and 0.102 mL Ph chlorocarbonate and stirred at room temp. for 2.5 h to give 330 mg N-[4-[6-(4-benzyloxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenyl]-N'-cyclopropylurea which (260 mg) was hydrogenolyzed over platinum oxide in ethanol overnight to give 160 mg N-[4-[6-(4-hydroxyphenyl)-7-(2-trimethylsilylethoxymethyl)-7H-pyrrolo[2,3-d]pyrimidin-4-yloxy]-2-chlorophenyl]-N'-cyclopropylurea (I). I showed IC₅₀ of 0.02 nM for inhibiting the vascular endothelial growth factor (VEGF)-stimulated sandwich tube formation in vascular endothelial cell.

IT **417714-38-6P**

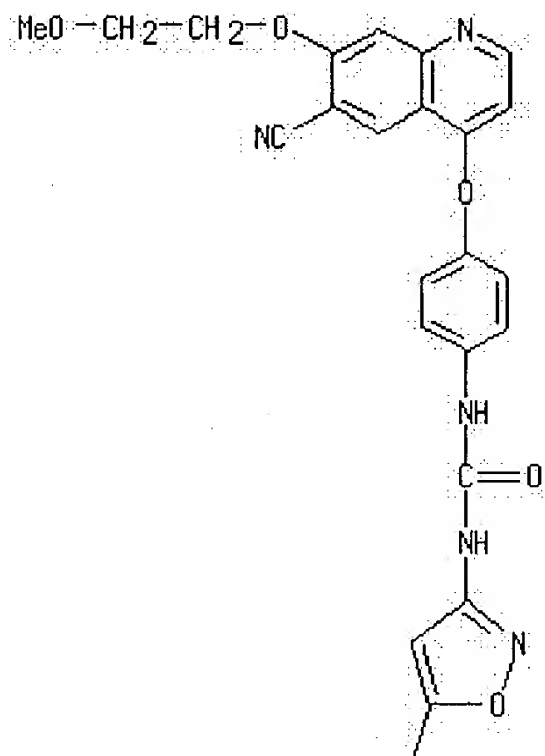
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of urea derivs. contg. nitrogenous arom. ring compds. as
angiogenesis inhibitors for prevention or treatment of diseases)

RN 417714-38-6 HCAPLUS

CN Urea, N-[4-[[6-cyano-7-(2-methoxyethoxy)-4-quinolinyl]oxy]phenyl]-N'-(5-methyl-3-isoxazolyl)- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file caold

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

94.80

416.29

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

TOTAL

ENTRY

SESSION

CA SUBSCRIBER PRICE

-7.00

-7.00

FILE 'CAOLD' ENTERED AT 10:51:51 ON 21 SEP 2004

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE COVERS 1907-1966

FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate
substance identification. Title keywords, authors, patent
assignees, and patent information, e.g., patent numbers, are
now searchable from 1907-1966. TIFF images of CA abstracts
printed between 1907-1966 are available in the PAGE

display formats.

This file supports REGISTRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

=> d his

(FILE 'HOME' ENTERED AT 10:32:23 ON 21 SEP 2004)

FILE 'REGISTRY' ENTERED AT 10:32:29 ON 21 SEP 2004

L1 STRUCTURE UPLOADED
L2 23 S L1
L3 377 S L1 FULL

FILE 'HCAPLUS' ENTERED AT 10:34:46 ON 21 SEP 2004

L4 34 S L3

FILE 'REGISTRY' ENTERED AT 10:36:32 ON 21 SEP 2004

L5 STRUCTURE UPLOADED
L6 8 S L5
L7 189 S L5 FULL

FILE 'HCAPLUS' ENTERED AT 10:39:55 ON 21 SEP 2004

L8 10 S L7
L9 3 S L8 AND DUMAS, J?/AU
L10 7 S L8 NOT L9
L11 0 S L10 AND KHIRE, U?/AU
L12 1 S L10 AND LOWINGER, T?/AU
L13 6 S L10 NOT L12
L14 0 S L13 AND PAULSEN, H?/AU
L15 0 S L13 AND RIEDL, B?/AU
L16 0 S L13 AND SCOTT, W?/AU
L17 0 S L13 AND SMITH, R?/AU
L18 0 S L13 AND WOOD, J?/AU
L19 0 S L13 AND HATOUM-MOKDAD, H?/AU
L20 0 S L13 AND JOHNSON, J?/AU
L21 0 S L13 AND LEE, W?/AU
L22 0 S L13 AND REDMAN, A?/AU
L23 0 S L13 AND SIBLEY, R?/AU
L24 0 S L13 AND RENICK, F?/AU
L25 0 S L13 AND RENICK, J?/AU

FILE 'CAOLD' ENTERED AT 10:51:51 ON 21 SEP 2004

=> s 17

L26 0 L7

=>